<u>AMENDMENTS TO THE CLAIMS:</u>

This listing of claims will replace all prior versions, and listings of claims in the application:

1. (Previously Amended) A compound of the formula:

Wherein the dashed bond represents a single or double bond;

Aryl signifies a monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;

 R^1 is H, OH, OC₁₋₃alkyl, C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;

 R^2 is H, halogen, C_{1-3} alkyl, $CONR^5R^6$, $S(=O)_mC_{1-3}$ alkyl, or C_{1-3} alkyl substituted optionally with OH, or OC_{1-3} alkyl; with the proviso that if Aryl is thiophene, then $R^2 \neq H$ or halo, and $R^1 \neq OH$

R³, R⁴ are independently H, C_{1.3}alkyl, or C_{1.3}alkyl substituted optionally with OH or OC_{1.3}alkyl:

R⁵, R⁶ are independently H, C₁₋₃alkyl, or C₂₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, where R⁵ and R⁶ optionally can be joined together to form a pyrrolidine or piperidine ring which can be either unsubstituted or substituted optionally with C₁₋₃alkyl, C₂₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃,

OC₁₋₃alkyl, or C₁₋₃alkyl;

n is 2 to 4; m is 0, 1 or 2

or a pharmaceutically acceptable salt or solvate thereof.

(Currently Amended) A compound of the formula:

Wherein the dashed bond represents a single or double bond;

Aryl signifies a fused phenyl or monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, prvimidine, pyridazine, and pyrazine;

- R¹ is H, C₁₋₅alkyl, C₃₋₅alkenyl, an aromatic ring selected from the group consisting of phenyl, thicnyl, pyridyl, and imidazoyl which is either unsubstituted or substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, halogen, or CF₃; or C₂₋₅alkyl substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl or an aromatic ring selected from the group consisting of phenyl, thienyl, pyridyl, and imidazoyl, which is either unsubstituted or substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, halogen, CF₃, S(=O)₂ NR⁵R⁶; or C₃₋₅alkenyl substituted optionally with OH, OC₁₋₃alkyl, or S(=O)_mC₁₋₃alkyl;
- R² is H. halogen, C₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, S(=O)₂ NR⁵R⁶, or C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;
- R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;
- R⁵, R⁶ are independently H, C₁₋₃alkyl, or C₂₋₃alkyl substituted optionally with OH, OC₁.

 3alkyl, where R⁵ and R⁶ optionally can be joined together to form a pyrrolidine or piperidine ring which can be either unsubstituted or substituted optionally with C₁₋₃alkyl, C₂₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;
- R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally

selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl,

n is 2 to 4; m is 0, 1 or 2 or a pharmaceutically acceptable salt or solvate thereof.

(Withdrawn) A compound of the formula:

 R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

R⁹ is phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄ alkyl;

R¹⁰ is C₁₋₄alkyl, or R¹⁰ can be joined to R⁹ to form a fused bicyclic ring system such as indoline;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

(Withdrawn) A Compound of the formula:

- R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;
- R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N. O. S. such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;
- R^{11} is C_{1-3} alkyl, phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C_{1-4} alkyl, halogen, OC_{1-4} alkyl;
- R¹² is C₁₋₄alkyl or a fused bicyclic heteroaromatic ring such as thieno[3,2-e]-1,2-thiazine, or 1,2-benzothiazine, or R¹² can be joined to R¹¹ to form a fused bicyclic ring system such as 2,3-dihydro-benzo[c]isoxazole;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

5. (Currently Amended) A method for lowering IOP which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

Wherein the dashed bond represents a single or double bond;

Aryl signifies a fused phenyl or monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;

R¹ is H, OH, OC₁₋₃alkyl, C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;

R² is H, halogen, C₁₋₃alkyl, CONR⁵R⁶, S(=O)_mC₁₋₃alkyl⁶, or C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;

R³, R⁴ are independently H, C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;

R⁵, R⁶ are independently H, C₁₋₃alkyl, or C₂₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, where R⁵ and R⁶ optionally can be joined together to form a pyrrolidine or piperidine ring which can be either unsubstituted or substituted optionally with C₁₋₃alkyl, C₂₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

n is 2 to 4;

m is 0, 1 or 2

or a pharmaceutically acceptable salt or solvate thereof.

6. (Currently Amended) A method for lowering IOP which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

Wherein the dashed bond represents a single or double bond;

- Aryl signifies a fused phenyl or monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;
- R¹ is H, C₁₋₅alkyl, C₃₋₅alkenyl, an aromatic ring selected from the group consisting of phenyl, thienyl, pyridyl, and imidazoyl, which is either unsubstituted or substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, halogen, CF₃, or S(=O)₂ NR⁵R⁶; or C₂₋₅alkyl substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl or an aromatic ring selected from the group consisting of phenyl, thienyl, pyridyl, and imidazoyl, which is either unsubstituted or substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, halogen, CF₃, S(=O)₂ NR⁵R⁶; or C₃₋₅alkenyl substituted optionally with OH, OC₁₋₃alkyl, or S(=O)_mC₁₋₃alkyl;
- R^2 is H, halogen, C_{1-3} alkyl, $S(=0)_m C_{1-3}$ alkyl, or C_{1-3} alkyl substituted optionally with OH, or OC_{1-3} alkyl;
- R³ & R⁴ are independently H, C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;
- R⁵, R⁶ are independently H, C₁₋₃alkyl, or C₂₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, where R⁵ and R⁶ optionally can be joined together to form a pyrrolidine or piperidine ring which can be either unsubstituted or substituted optionally with C₁₋₃alkyl, C₂₋₃alkyl substituted optionally with OH or OC₁.

 3alkyl;
- R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁₋₃

3alkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

n is 2 to 4; m is 0, 1 or 2 or a pharmaceutically acceptable salt or solvate thereof.

7. (Withdrawn) A method for lowering IOP which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

 \mathbb{R}^3 & \mathbb{R}^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

R⁹ is phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄alkyl;

R¹⁰ is C₁₋₄alkyl, or R¹⁰ can be joined to R⁹ to form a fused bicyclic ring system such as indoline;

n is 2 to 4 and any pharmaceutically acceptable salts and solvates.

8. (Withdrawn) A method for lowering IOP which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

- R³ & R⁴ are independently H, C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;
- R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;
 - R^{11} is C_{1-3} alkyl, phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C_{1-4} alkyl, halogen, OC_{1-4} alkyl;
 - R^{12} is $C_{1.48}$ lkyl or a fused bicyclic heteroaromatic ring such as thieno[3,2-e]-1,2-thiazine, or 1,2-benzothiazine, or R^{12} can be joined to R^{11} to form a fused bicyclic ring system such as 2,3-dihydro-benzo[c]isoxazole;

n is 2 to 4 and any pharmaceutically acceptable salts and solvates.

9. (Currently Amended) A method for improving blood flow to the optic nerve head and the retina which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

Wherein the dashed bond represents a single or double bond;

Aryl signifies a fused phenyl or monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;

 R^1 is H, OH, OC₁₋₃alkyl, C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;

 R^2 is H, halogen, C_{1-3} alkyl, $CONR^5R^6$, $S(=O)_mC_{1-3}$ alkyl, or C_{1-3} alkyl substituted optionally with OH, or OC_{1-3} alkyl;

 \mathbb{R}^3 , \mathbb{R}^4 are independently H, \mathbb{C}_{1-3} alkyl, or \mathbb{C}_{1-3} alkyl substituted optionally with OH or \mathbb{OC}_{1-3} alkyl;

R⁵, R⁶ are independently H, C₁₋₃alkyl, or C₂₋₃alkyl substituted optionally with OH, OC₁.

3alkyl, where R⁵ and R⁶ optionally can be joined together to form pyrrolidine or piperidine ring which can be either unsubstituted or substituted optionally with C₁₋₃alkyl, C₂₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

n is 2 to 4;

m is 0, 1 or 2

or a pharmaceutically acceptable salt or solvate thereof.

10. (Currently Amended) A method for improving blood flow to the optic nerve head and the retina which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

Wherein the dashed bond represents a single or double bond;

Aryl signifies a fused phenyl or monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;

- R¹ is H, C₁₋₅alkyl, C₃₋₅alkenyl, an aromatic ring selected from the group consisting of phenyl, thienyl, pyridyl, and imidazoyl, which is either unsubstituted or substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, halogen, CF₃, or S(=O)₂ NR⁵R⁶; or C₂₋₅alkyl substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl or an aromatic ring selected from the group consisting of phenyl, thienyl, pyridyl, and imidazoyl, which is either unsubstituted or substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, halogen, CF₃, S(=O)₂ NR⁵R⁶; or C₃₋₅alkenyl substituted optionally with OH, OC₁₋₃alkyl, or S(=O)_mC₁₋₃alkyl;
- R² is H, halogen, C₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;
- R³ & R⁴ are independently H, C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;
- R⁵, R⁶ are independently H, C₁₋₃alkyl, or C₂₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, where R⁵ and R⁶ optionally can be joined to form a pyrrolidine or piperidine ring which can be either unsubstituted or substituted optionally with C₁₋₃alkyl, C₂₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;
- R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁.

³alkyl, or phenyl which can be unsubstituted or substituted optionally with balogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with balogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

n is 2 to 4; m is 0, 1 or 2 or a pharmaceutically acceptable salt or solvate thereof.

11. (Withdrawn) A method for improving blood flow to the optic nerve head and the retina which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

 R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

R⁹ is phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄ alkyl;

R¹⁰ is C₁₋₄alkyl, or R¹⁰ can be joined to R⁹ to form a fused bicyclic ring system such as

indoline;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

12. (Withdrawn) A method for improving blood flow to the optic nerve head and the retina which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

 R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

R¹¹ is C₁₋₃alkyl, phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄alkyl;

R¹² is C₁₋₄alkyl or a fused bicyclic heteroaromatic ring such as thieno[3,2-e]-1,2-thiazine, or 1,2-benzothiazine, or R¹² can be joined to R¹¹ to form a fused bicyclic ring system such as 2,3-dihydro-benzo[c]isoxazole;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

13. (Currently Amended) A method for treating retinal diseases selected from the group consisting of glaucoma, age related macular degeneration (ARMD), optic neuritis, ischemic disorders, diabetic retinopathy, and retinal edema which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

Wherein the dashed bond represents a single or double bond;

Aryl signifies a fused phenyl or monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;

R¹ is H, OH, OC₁₋₃alkyl, C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;

R² is H, halogen, C₁₋₃alkyl, CONR⁵R⁶, S(=O)_mC₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;

R³, R⁴ are independently H, C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;

R⁵, R⁶ are independently H, C₁₋₃alkyl, or C₂₋₃alkyl substituted optionally with OH, OC₁.

3alkyl, where R⁵ and R⁶ optionally can be joined together to form a pyrrolidine or piperidine ring which can be either unsubstituted or substituted optionally with C₁₋₃alkyl, C₂₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃,

OC1-3alkyl, or C1-3alkyl;

n is 2 to 4;

m is 0, 1 or 2

or a pharmaceutically acceptable salt or solvate thereof.

14. (Currently Amended) A method for treating retinal diseases selected from the group consisting of glaucoma, age related macular degeneration (ARMD), optic neuritis, ischemic disorders, diabetic retinopathy, and retinal edema which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

Wherein the dashed bond represents a single or double bond;

- Aryl signifies a fused phenyl or monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;
- R¹ is H, C₁₋₅alkyl, C₃₋₅alkenyl, an aromatic ring selected from the group econsisting consisting of phenyl, thienyl, pyridyl, and imidazoyl, which is either unsubstituted or substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, halogen, CF₃, or S(=O)₂ NR⁵R⁶; or C₂₋₅alkyl substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl or an aromatic ring selected from the group consisting of phenyl, thienyl, pyridyl, and imidazoyl which is either unsubstituted or substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, halogen, CF₃, S(=O)₂ NR⁵R⁶; or C₃₋₅alkenyl substituted optionally with OH, OC₁₋₃alkyl, or S(=O)_mC₁₋₃alkyl;
- R² is H, balogen, C₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;
- R³ & R⁴ are independently H, C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;
- R⁵, R⁶ are independently H, C₁₋₃alkyl, or C₂₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, where R⁵ and R⁶ optionally can be joined together to form a pyrrolidine or piperidine ring which can be either unsubstituted or substituted

optionally with C_{1-3} alkyl, C_{2-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

n is 2 to 4; m is 0, 1 or 2 or a pharmaceutically acceptable salt or solvate thereof.

15. (Withdrawn) A method for treating retinal diseases which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

$$R^{10}$$
 $(CR^3R^4)_n$
 R^7

 R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl,

or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁. ₃alkyl;

R⁹ is phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄alkyl;

R¹⁰ is C₁₋₄alkyl, or R¹⁰ can be joined to R⁹ to form a fused bicyclic ring system such as indoline;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

16. (Withdrawn) A method for treating retinal diseases which comprises administering to a person in need thereof, a composition comprising an effective amount of a compound of the formula:

 R^3 & R^4 are independently H, $C_{1\text{--}3}$ alkyl, or $C_{1\text{--}3}$ alkyl substituted optionally with OH or $OC_{1\text{--}3}$ alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidem, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁

3alkyl;

R¹¹ is C₁₋₃alkyl, phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C1-4 alkyl, halogen, OC1-4alkyl;

 \mathbb{R}^{12} is $C_{1,4}$ alkyl or a fused bicyclic heteroaromatic ring such as thieno[3,2-e]-1,2-thiazine, or 1,2-benzothiazine, or R12 can be joined to R11 to form a fused bicyclic ring system such as 2,3-dihydro-benzo[c]isoxazole;

n is 2 to 4

and any pharmaceutically acceptable salts and solvales.

- (Cancelled) 17.
- (Cancelled) 18.
- (Withdrawn) A composition for lowering IOP comprising a pharmaceutically 19. effective amount of a compound of the formula:

- R3 & R4 are independently H, C1.3alkyl, or C1.3alkyl substituted optionally with OH or OC1-3alkyl;
- R7, R8 are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ^3 -piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C1-3alkyl, C1-3alkyl substituted optionally with OH, OC1-3alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF3, OC1.3alkyl, or C1-3alkyl, or substituted on nitrogen with C1-4alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃ 3alkyl;

R⁹ is phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄ alkyl;

R¹⁰ is C₁₋₄alkyl, or R¹⁰ can be joined to R⁹ to form a fused bicyclic ring system such as indoline;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

20. (Withdrawn) A composition for lowering IOP comprising a pharmaceutically effective amount of a compound of the formula:

- R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;
- R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;
 - R¹¹ is C₁₋₃alkyl, phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄alkyl;
 - R¹² is C₁₋₄alkyl or a fused bicyclic heteroaromatic ring such as thieno[3,2-e]-1,2-thiazine, or 1,2-benzothiazine, or R¹² can be joined to R¹¹ to form a fused bicyclic ring system such as 2,3-dihydro-benzo[c]isoxazole;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

- 21. (Cancelled)
- 22. (Cancelled)
- 23. (Withdrawn) A composition for improving blood flow to the optic nerve head and the retina comprising a pharmaceutically effective amount of a compound of the formula:

$$R^{10}$$
 $(CR^3R^4)_n$
 R^7

- R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;
- R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;
- R⁹ is phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄alkyl;
- R¹⁰ is C₁₋₄alkyl, or R¹⁰ can be joined to R⁹ to form a fused bicyclic ring system such as indoline;

n is 2 to 4

and any pharmaccutically acceptable salts and solvates.

24. (Withdrawn) A composition for improving blood flow to the optic nerve head and the retina comprising a pharmaceutically effective amount of a Compound of the formula:

 R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

R¹¹ is C₁₋₃alkyl, phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄alkyl;

R¹² is C₁₋₄alkyl or a fused bicyclic heteroaromatic ring such as thieno[3,2-e]-1,2-thiazine, or 1,2-benzothiazine, or R¹² can be joined to R¹¹ to form a fused bicyclic ring system such as 2,3-dihydro-benzo[c]isoxazole;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

25. (Cancelled)

26. (Cancelled)

27. (Withdrawn) A composition for treating retinal diseases comprising a pharmaceutically effective amount of a compound of the formula:

 R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

R⁹ is phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄ alkyl;

R¹⁰ is C₁₋₄alkyl, or R¹⁰ can be joined to R⁹ to form a fused bicyclic ring system such as indoline;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

28. (Withdrawn) A composition for treating retinal diseases comprising a pharmaceutically effective amount of a compound of the formula:

 R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

R¹¹ is C₁₋₃alkyl, phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄alkyl, halogen, OC₁₋₄alkyl;

R¹² is C₁₋₄alkyl or a fused bicyclic heteroaromatic ring such as thieno[3,2-e]-1,2-thiazine, or 1,2-benzothiazine, or R¹² can be joined to R¹¹ to form a fused bicyclic ring system such as 2,3-dihydro-benzo[c]isoxazole;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

29. (Withdrawn) A method for improving blood flow to the optic nerve head or the retina which comprises administering to a person in need thereof, a composition comprising a pharmaceutically effective amount of a compound with 5HT₇ receptor affinity.

- 30. (Withdrawn) A composition for improving blood flow to the optic nerve head or the retina comprising a pharmaceutically effective amount of a compound with 5HT₇ receptor affinity.
- 31. (Withdrawn) A method for providing neuroprotection to the optic nerve head or the retina which comprises administering to a person in need thereof, a composition comprising a pharmaceutically effective amount of a compound with 5HT₇ receptor affinity.
- 32. (Withdrawn) A composition for providing neuroprotection to the optic nerve head or the retina comprising a pharmaceutically effective amount of a compound with 5HT₇ receptor affinity.
- 33. (Withdrawn) A method for treating retinal diseases which comprises administering to a person in need thereof, a composition comprising a pharmaceutically effective amount of a compound with 5HT₇ receptor affinity.
- 34. (Cancelled)
- 35. (Withdrawn) A composition for treating retinal diseases comprising a pharmaceutically effective amount of a compound with 5HT₇ receptor affinity.
- 36. (Withdrawn) The composition of Claim 35 wherein the retinal diseases are selected from the group consisting of glaucoma, age related macular degeneration, optic neuritis, ischemic disorders, diabetic retinopathy, and retinal edema.
- 37. (Withdrawn) A method for lowering IOP which comprises administering to a person in need thereof, a composition comprising a pharmaceutically effective amount of a compound with 5HT₇ receptor affinity.
- 38. (Withdrawn) A composition for lowering IOP comprising a pharmaceutically effective amount of a compound with 5HT₂ receptor affinity.

39. (Currently Amended) A method for treating persons suffering from a sleeping disorder, depression, schizophrenia, anxiety, circadian rhythm disorders, and centrally and peripherally mediated hypertension, which comprises, administering a composition comprising a pharmaceutically effective amount of a compound of the formula:

Wherein the dashed bond represents a single or double bond

Aryl signifies a fused phenyl or monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;

R¹ is H, OH, OC₁₋₃alkyl, C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;

 R^2 is H, halogen, C_{1-3} alkyl, $CONR^5R^6$, $S(=O)_mC_{1-3}$ alkyl, $S(=O)_2$ NR^5R^6 , C_{1-3} alkyl substituted optionally with OH, or OC_{1-3} alkyl;

R³, R⁴ are independently H, C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;

R⁵, R⁶ are independently H, C_{1.3}alkyl, or C_{2.3}alkyl substituted optionally with OH, OC₁.

3alkyl, where R⁵ and R⁶ optionally can be joined together to form a pyrrolidine or piperidine ring which can be either unsubstituted or substituted optionally with C_{1.3}alkyl, C_{2.3}alkyl substituted optionally with OH or OC_{1.3}alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ¹-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

n is 2 to 4;

m is 0, 1 or 2 or a pharmaceutically acceptable salt or solvate thereof.

40. (Currently Amended) A method for treating persons suffering from a sleeping disorder, depression, schizophrenia, anxiety, obsessive compulsive disorder, circadian rhythem disorders, and centrally and peripherally mediated hypertension which comprises, administering a composition comprising a pharmaceutically effective amount of a compound of the formula:

Wherein the dashed bond represents a single or double bond;

Aryl signifies a fused phenyl or monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;

- R¹ is H, C₁₋₅alkyl, C₃₋₅alkenyl, an aromatic ring selected from the group consisting of phenyl, thienyl, pyridyl, and imidazoyl, which is either unsubstituted or substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, halogen, CF₃, or S(=O)₂ NR⁵R⁶; or C₂₋₅alkyl substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl or an aromatic ring selected from the group consisting of phenyl, thienyl, pyridyl, and imidazoyl which is either unsubstituted or substituted optionally with OH, OC₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, halogen, CF₃, S(=O)₂ NR⁵R⁶; or C₃₋₅alkenyl substituted optionally with OH, OC₁₋₃alkyl, or S(=O)_mC₁₋₃alkyl;
- R^2 is H, halogen, C_{1-3} alkyl, $S(=O)_mC_{1-3}$ alkyl, $S(=O)_2$ NR^5 R^6 , or C_{1-3} alkyl substituted optionally with OH, or OC_{1-3} alkyl;
- R³ & R⁴ are independently H, C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;
- R⁵, R⁶ are independently H, C₁₋₃alkyl, or C₂₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, where R⁵ and R⁶ optionally can be joined together to form a pyrrolidine or piperidine and which can be either unsubstituted or substituted optionally with C₁₋₃alkyl, C₂₋₃alkyl substituted optionally with OH or OC₁₋₃

3alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ^3 -piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C1-3alkyl, or C1-3alkyl substituted optionally with OH, OC1-3alkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋ 4alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF3, OC1-3alkyl, or C1-3alkyl;

n is 2 to 4; m is 0, 1 or 2 or a pharmaceutically acceptable salt or solvate thereof.

(Withdrawn) A method for treating persons suffering from a sleeping disorder, 41. depression, schizophrenia, anxiety, obsessive compulsive disorders, circadian rhythm disorders, and centrally and peripherally mediated hypertension which comprises, administering a composition comprising a pharmaceutically effective amount of a compound of the formula:

R3 & R4 are independently H, C1-3alkyl, or C1-3alkyl substituted optionally with OH or OC₁-3alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ^3 -piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which (

can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

R⁹ is phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, halogen, OC₁₋₄ alkyl;

R¹⁰ is C₁₋₄alkyl, or R¹⁰ can be joined to R⁹ to form a fused bicyclic ring system such as indoline;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates.

42. (Withdrawn) A method for treating persons suffering from a sleeping disorder, depression, schizophrenia, anxiety, obsessive compulsive disorder, circadian rhythm disorders, and centrally and peripherally mediated hypertension which comprises, administering a composition comprising a pharmaceutically effective amount of a compound of the formula:

$$R^{11} \underbrace{N}_{N} S - R^{12}$$

$$(CR^{3}R^{4})_{n} - N$$

 R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl,

or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

R¹¹ is C₁₋₃alkyl, phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄alkyl, halogen, OC₁₋₄alkyl;

R¹² is C₁₋₄alkyl or a fused bicyclic heteroaromatic ring such as thieno[3,2-e]-1,2-thiazine, or 1,2-benzothiazine, or R¹² can be joined to R¹¹ to form a fused bicyclic ring system such as 2,3-dihydro-benzo[c]isoxazole;

n is 2 to 4

and any pharmaccutically acceptable salts and solvates.

43. (Previously Amended) A pharmaceutical composition comprising a pharmaceutically effective amount of a compound of the formula:

Wherein the dashed bond represents a single or double bond;

Aryl signifies a monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;

R¹ is H, OH, OC₁₋₃alkyl, C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, or OC₁₋₃alkyl;

 R^2 is H, halogen, C_{1-3} alkyl, $CONR^5R^6$, $S(=O)_mC_{1-3}$ alkyl, or C_{1-3} alkyl substituted optionally with OH, or OC_{1-3} alkyl; with the proviso that if Aryl is thiophene, then $R^2 \neq H$ or halo, and $R^1 \neq OH$

R³, R⁴ are independently H, C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;

R⁵, R⁶ are independently H, C_{1:3}alkyl, or C₂₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, where R⁵ and R⁶ optionally can be joined together to form a pyrrolidine or piperidine ring which can be either unsubstituted or substituted optionally with C₁₋₃alkyl, C₂₋₃alkyl substituted optionally with OH or OC₁₋₃alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, Δ^3 -piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF3, OC1-3alkyl, or C1-3alkyl;

n is 2 to 4;

m is 0, 1 or 2

or a pharmaceutically acceptable salt or solvate thereof in a pharmaceutically acceptable carrier.

A pharmaceutical composition comprising a (Previously Amended) 44. pharmaceutically effective amount of a compound of the formula:

Wherein the dashed bond represents a single or double bond;

Aryl signifies a monocyclic heteroaromatic ring selected from the group consisting of thiophene, furan, pyrrole, pyridine, pryimidine, pyridazine, and pyrazine;

R1 is H, C1.5alkyl, C3.5alkenyl, an aromatic ring selected from the group consisting of phenyl, thienyl, pyridyl, and imidazoyl which is either unsubstituted or substituted optionally with OH, OC1-3alkyl, S(=O)mC1-3alkyl, halogen, or_CF3; or C2.5alkyl substituted optionally with OH, OC1-3alkyl, S(=O)mC1-3alkyl or an aromatic ring selected from the group consisting of phenyl, thienyl, pyridyl, and imidazoyl, which is either unsubstituted or substituted optionally with OH, OC_{1-3} alkyl, $S(=O)_mC_{1-3}$ alkyl, halogen, CF_3 , $S(=O)_2$ NR^5R^6 ; or C_{3-5} alkenyl substituted optionally with OH, OC1.3alkyl, or S(=O)mC1.3alkyl;

R² is H, halogen, C₁₋₃alkyl, S(=O)_mC₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, or OC1.3alkyl;

R3 & R4 are independently H, C1-3alkyl, or C1-3alkyl substituted optionally with OH or

OC1-3alkyl;

- R^5 , R^6 are independently H, $C_{1\cdot 3}$ alkyl, or $C_{2\cdot 3}$ alkyl substituted optionally with OH, $OC_{1\cdot 3}$ 3alkyl, where R5 and R6 optionally can be joined together to form a pyrrolidine or piperidine ring which can be either unsubstituted or substituted optionally with C_{1-3} alkyl, C_{2-3} alkyl substituted optionally with OH or OC_{1-1}
- R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring selected from the group consisting of pytrolidine, piperidine, Δ^3 -piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, or C₁₋₃alkyl substituted optionally with OH, OC₁salkyl, or phenyl which can be unsubstituted or substituted optionally with halogen, CF3, OC1-3alkyl, or C1-3alkyl, or substituted on nitrogen with C1-4alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF3, OC1-3alkyl, or C1-3alkyl;

n is 2 to 4;

m is 0, 1 or 2

or a pharmaceutically acceptable salt or solvate thereof in a pharmaceutically acceptable carrier.

(Withdrawn) A composition comprising a pharmaceutically effective amount of a 45. compound of the formula:

R3 & R4 are independently H, C1.3alkyl, or C1.3alkyl substituted optionally with OH or OC1-3alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ^3 -piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from

C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl;

R⁹ is phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C₁₋₄ alkyl, balogen, OC₁₋₄ alkyl;

R¹⁰ is C₁₋₄alkyl, or R¹⁰ can be joined to R⁹ to form a fused bicyclic ring system such as indoline;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates in a pharmaceutically acceptable carrier.

46. (Withdrawn) A composition comprising a pharmaceutically effective amount of a compound of the formula:

 R^3 & R^4 are independently H, C_{1-3} alkyl, or C_{1-3} alkyl substituted optionally with OH or OC_{1-3} alkyl;

R⁷, R⁸ are together with the nitrogen atom to which they are attached incorporated into a heterocyclic ring of 5 to 8 atoms which may include a second heteroatom selected from N, O, S, such as pyrrolidine, piperidine, Δ³-piperidein, piperazine, morpholine or thiomorpholine which can be unsubstituted or substituted on carbon with one or more substituents optionally selected from C₁₋₃alkyl, C₁₋₃alkyl substituted optionally with OH, OC₁₋₃alkyl, phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or substituted on nitrogen with C₁₋₄alkoxy or phenyl which can be unsubstituted or substituted optionally with halogen, CF₃, OC₁₋₃alkyl, or C₁₋₃alkyl, or C

3elkyl;

- R^{11} is C_{1-3} alkyl, phenyl or a monocyclic heteroaromatic ring which can be unsubstituted or substituted with C1-4 alkyl, halogen, OC1-4 alkyl;
- R^{12} is C_{1-4} alkyl or a fused bicyclic heteroaromatic ring such as thieno[3,2-e]-1,2-thiazine, or 1,2-benzothiazine, or R12 can be joined to R11 to form a fused bicyclic ring system such as 2,3-dihydro-benzo[c]isoxazole;

n is 2 to 4

and any pharmaceutically acceptable salts and solvates in a pharmaceutically acceptable

- (Original) The Compound of Claim 1 selected from the group consisting of: 47. 6-Chloro-2-[4-[4-(2H-benzimidazo-2-oxo-1-yl)piperidin-1-yl]butyl]-2Hthieno[3,2-e]-1,2-thiazine 1,1-dioxide;
 - 6-Chloro-2-[4-(4-phenylpiperazin-1-yl)butyl]-2H-thieno[3,2-e]-1,2-thiazine 1,1dioxide;
 - 6-Chloro-2-[4-[4-(2-fluorophenyl)piperazin-1-yl]butyl]-2H-thieno[3,2-e]-1,2thiazine 1,1-dioxide;
 - 6-Chloro-2-[3-[4-(3-trifluoromethylphenyl)pipcrazin-1-yl]propyl]-2H-thieno[3,2e]-1,2-thiazine 1,1-dioxide;
 - 6-Chloro-2-[3-[4-(2H-benzimidazol-2-oxo)piperidin-1-yl]propyl]-2H-thieno[3,2e]-1,2-thiazine 1,1-dioxide.
- (Withdrawn) The Compound of Claim 3 selected from the group consisting of: 48.
- 3-[4-(3-Chlorophenyl)piperazin-1-yl]propylsulfonyl-2, 3-dihydro-1 H-indole;
- 3-(1,2,3,4-Tetrabydroisoquinolin-2-yl)propylsulfonyl-2,3-dihydro-1H-indole;
- 3-[4-(3-Trifluoromethylphenyl)piperazin-1-yl]propylsulfonyl-2,3-dihydro-1H-indole;
- 3-[4-(2-Methoxyphenyl)piperazin-1-yl]propylsulfonyl-2,3-dihydro-1H-indole;
- 3-(1,2,3,4-Tetrahydroisoquinolin-2-yl)-N-methyl-N-phcnyl-propylsulfonamide;

- 49. (Withdrawn) The Compound of Claim 4 selected from the group consisting of:
- N-[3-[4-(3-Chlorophenyl)piperazin-1-yl]propyl]-N-(4-methoxyphenyl)-propanesulfonamide;
- N-[3-(1,2,3,4-Tetrahydroisoquinolin-2-yl)propyl]-N-(4-methoxyphenyl)-propanesulfonamide;
- N-[3-[4-(3-Chlorophenyl)piperazin-1-yl]propyl]-N-(4-methoxyphenyl)-propanesulfonamide;
- N-[3-[4-(2-Methoxyphenyl)piperazin-1-yl]propyl]-N-(4-methoxyphenyl)-propanesulfonamide;
- $\label{eq:N-3-4-4-methoxyphenyl} $$N-[3-[4-(2-Chlorophenyl)piperazin-1-yl]propyl]-N-(4-methoxyphenyl)-propanesulfonamide.$

This Page is Inserted by IFW Indexing and Scanning Operations and is not part of the Official Record

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images include but are not limited to the items checked:

☐ BLACK BORDERS
☐ IMAGE CUT OFF AT TOP, BOTTOM OR SIDES
☐ FADED TEXT OR DRAWING
☐ BLURRED OR ILLEGIBLE TEXT OR DRAWING
☐ SKEWED/SLANTED IMAGES
☐ COLOR OR BLACK AND WHITE PHOTOGRAPHS
☐ GRAY SCALE DOCUMENTS
☐ LINES OR MARKS ON ORIGINAL DOCUMENT
☐ REFERENCE(S) OR EXHIBIT(S) SUBMITTED ARE POOR QUALITY
□ OTHER:

IMAGES ARE BEST AVAILABLE COPY.

As rescanning these documents will not correct the image problems checked, please do not report these problems to the IFW Image Problem Mailbox.